

Applicant : Anders Brilsson et al.
 Serial No. : 10/593,543
 Filed : September 20, 2006
 Page : 2 of 24

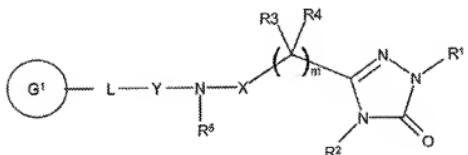
Attorney's Docket No.: 06275-522US1 / 101414-1P US

Amendments to the Claims:

This listing of claims replaces all prior versions and listings of claims in the application:

Listing of Claims:

1. (Currently Amended) A compound of formula (I) or a pharmaceutically acceptable salt thereof



(I)

wherein

R¹ and R² independently represent H or C1 to 6 alkyl; said alkyl being optionally further substituted by an aryl ring or an aromatic heterocyclic ring containing 1 to 3 heteroatoms independently selected from O, S and N; said aromatic ring being optionally further substituted by halogen, CF₃, C1 to 4 alkyl or C1 to 4 alkoxy;

Author Search

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FILE COVERS 1907 - 2 Oct 2009 VOL 151 ISS 15
 FILE LAST UPDATED: 1 Oct 2009 (20091001/ED)
 REVISED CLASS FIELDS (/NCL) LAST RELOADED: Aug 2009
 USPTO MANUAL OF CLASSIFICATIONS THESAURUS ISSUE DATE: Aug 2009

HCplus now includes complete International Patent Classification (IPC) reclassification data for the third quarter of 2009.

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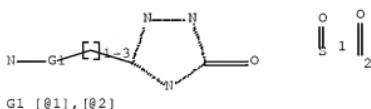
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| L9 | 535 | SEA FILE=HCPLUS SPE=ON ABB=ON PLU=ON ERIKSSON A?/AU |
| L10 | 10 | SEA FILE=HCPLUS SPE=ON ABB=ON PLU=ON LEPISTO M?/AU |

L11 1 SEA FILE=HCAPLUS SPE=ON ABB=ON PLU=ON (L10 OR L9) AND L8

=> FILE WPIX
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FILE LAST UPDATED: 28 SEP 2009 <20090928/UP>
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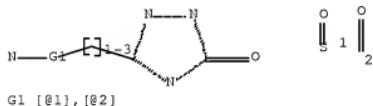
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G1 [01], [02]

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PROCESSING COMPLETED FOR L11

PROCESSING COMPLETED FOR L15

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ANSWER '1' FROM FILE HCPLUS

=> D IBIB ED ABS HITSTR L33 1

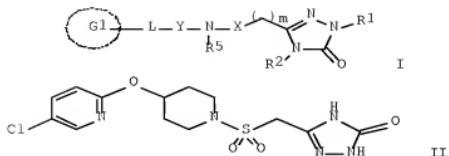
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 ACCESSION NUMBER: 2005:1106854 HCPLUS Full-text
 DOCUMENT NUMBER: 143:387043
 TITLE: Preparation of triazolone derivatives as MMP
 inhibitors for the treatment of asthma
 INVENTOR(S): Eriksson, Anders; Lepistoe, Matti
 PATENT ASSIGNEE(S): AstraZeneca AB, Swed.
 SOURCE: PCT Int. Appl., 53 pp.
 CODEN: PIXXD2
 DOCUMENT TYPE: Patent
 LANGUAGE: English
 FAMILY ACC. NUM. COUNT: 1
 PATENT INFORMATION:

| PATENT NO. | KIND | DATE | APPLICATION NO. | DATE |
|--|------|----------|------------------|------------|
| WO 2005095362 | A1 | 20051013 | WO 2005-SE448 | 20050329 |
| W: AB, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BW, BY, BZ, CA, CH, CN, CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, EG, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NA, NI, NO, NZ, OM, PG, PH, PL, PT, RO, RU, SC, SD, SE, SG, SK, SL, SM, SY, TJ, TM, TN, TR, TT, TZ, UA, UG, US, UZ, VC, VN, YU, ZA, ZM, ZW | | | | |
| RW: BW, GH, GM, KE, LS, MW, MZ, NA, SD, SL, SZ, TZ, UG, ZM, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM, AT, BE, BG, CH, CY, CZ, DE, DK, EE, ES, FI, FR, GB, GR, HU, IE, IS, IT, LT, LU, MC, NL, PL, PT, RO, SE, SI, SK, TR, BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN, TD, TG | | | | |
| EP 1732903 | A1 | 20061220 | EP 2005-722275 | 20050329 |
| EP 1732903 | B1 | 20090218 | | |
| R: AT, BE, BG, CH, CY, CZ, DE, DK, EE, ES, FI, FR, GB, GR, HU, IE, IS, IT, LI, LT, LU, MC, NL, PL, PT, RO, SE, SI, SK, TR | | | | |
| CN 1960979 | A | 20070509 | CN 2005-80017672 | 20050329 |
| JP 2007530672 | T | 20071101 | JP 2007-506108 | 20050329 |
| AT 423105 | T | 20090315 | AT 2005-722275 | 20050329 |
| ES 2320679 | T3 | 20090527 | ES 2005-722275 | 20050329 |
| US 20070219217 | A1 | 20070920 | US 2006-593543 | 20060920 |
| IN 2006DN05541 | A | 20070803 | IN 2006-DNS541 | 20060922 |
| HK 1099751 | A1 | 20090508 | HK 2007-105624 | 20070529 |
| PRIORITY APPLN. INFO.: | | | SE 2004-850 | A 20040330 |
| | | | WO 2005-SE448 | W 20050329 |

OTHER SOURCE(S): CASREACT 143:387043; MARPAT 143:387043

ED Entered STN: 14 Oct 2005

GI



AB Title compds. represented by the formula I [wherein R1, R2 = independently H, Cl or (un)substituted alkyl; R3, R4 = independently H, Cl, (un)substituted alkyl or R3R4 = (hetero)cyclyl; m = 1-3; X = SO, SO2 or CO; R5 = H, Cl or (un)substituted alkyl; Y = a direct bond or NR5Y = azacyclic ring; L = a direct bond, O, amino, etc.; G1 = (un)substituted cyclic ring; and pharmaceutically acceptable salts or solvates thereof] were prepared as metalloproteinase (MMP) inhibitors. For example, II was provided in a multi-step synthesis starting from the reaction of 5-(chloromethyl)-2,4-dihydro-3H-1,2,4-triazol-3-one with benzyl mercaptan. I were tested for inhibition of human MMP12, MMP9, MMP2, MMP19, MMP14 and MMP8. I and their pharmaceutical compns. are useful as MMP inhibitors for the treatment of asthma or other MMP-12 and/or MMP-9 mediated diseases (no data).

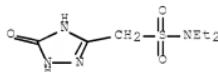
IT 866602-62-2P, N,N-Diethyl-1-(5-oxo-4,5-dihydro-1H-1,2,4-triazol-3-yl)methanesulfonamide

RL: BYP (Byproduct); PREP (Preparation)

(preparation of triazolone derivs. as MMP inhibitors for treatment of asthma)

RN 866602-62-2 HCPLUS

CN 1H-1,2,4-Triazole-3-methanesulfonamide, N,N-diethyl-2,5-dihydro-5-oxo-(CA INDEX NAME)



IT 866602-59-7P, 5-[[4-[(5-Chloropyridin-2-yl)oxy]piperidin-1-yl]sulfonyl]methyl]-2,4-dihydro-3H-1,2,4-triazol-3-one
 866602-63-3P, 5-[2-[[4-[(5-Chloropyridin-2-yl)oxy]piperidin-1-yl]sulfonyl]ethyl]-2,4-dihydro-3H-1,2,4-triazol-3-one
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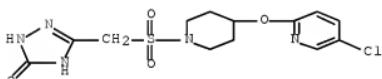
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 866602-76-8P, N-Benzyl-1-(5-oxo-4,5-dihydro-1H-1,2,4-triazol-3-yl)methanesulfonamide 866602-77-9P,
 1-(5-Oxo-4,5-dihydro-1H-1,2,4-triazol-3-yl)-N-(2-phenylethyl)methanesulfonamide 866602-78-0P,
 5-[2-[(4-Chlorophenyl)piperidin-1-yl]sulfonyl]ethyl]-2,4-dihydro-3H-1,2,4-triazol-3-one 866602-79-1P,
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 5-[3-[(4-Chlorophenyl)piperidin-1-yl]sulfonyl]propyl]-2,4-dihydro-3H-1,2,4-triazol-3-one 866602-81-5P,
 5-[3-[(4-Chlorophenyl)piperazin-1-yl]sulfonyl]propyl]-2,4-dihydro-3H-1,2,4-triazol-3-one

RL: PAC (Pharmacological activity); SPN (Synthetic preparation); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); USES (Uses)

(preparation of triazolone derivs. as MMP inhibitors for treatment of asthma)

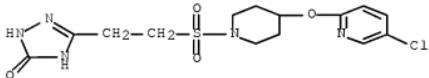
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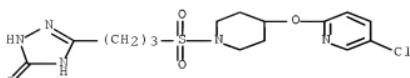
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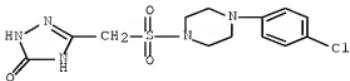


RN 866602-67-7 HCPLUS

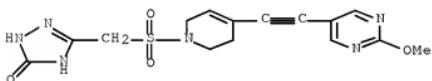
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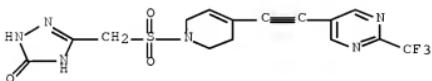
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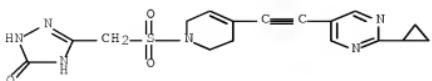
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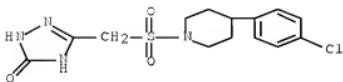
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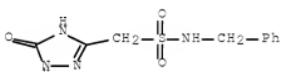
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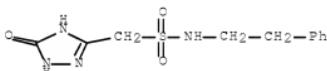
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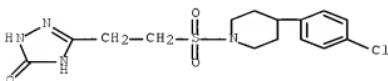
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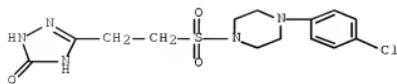
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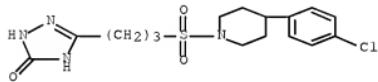


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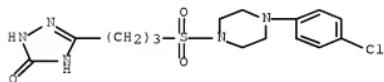
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RN 866602-81-5 HCPLUS

CN 3H-1,2,4-Triazol-3-one, 5-[3-[(4-(4-chlorophenyl)-1-piperazinyl)sulfonyl]propyl]-1,2-dihydro- (CA INDEX NAME)



REFERENCE COUNT:

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REVISED CLASS FIELDS (NCL) LAST RELOADED: Aug 2009
USPTO MANUAL OF CLASSIFICATIONS THESAURUS ISSUE DATE: Aug 2009

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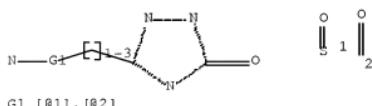
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 MOST RECENT UPDATE: 200962 <200962/DW>
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>>> IPC, ECLA, US National Classifications and Japanese F-Terms
 and FI-Terms have been updated with reclassifications to
 mid-June 2009.
 No update date (UP) has been created for the reclassified
 documents, but they can be identified by
 specific update codes (see HELP CLA for details)<<<

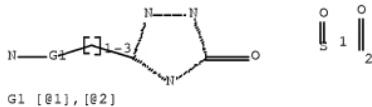
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=> FILE BEILSTEIN
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FILE LAST UPDATED ON May 17, 2009

FILE COVERS 1779 TO 2008.

*** FILE CONTAINS 10,593,281 SUBSTANCES ***

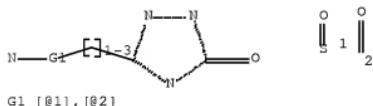
>>> PLEASE NOTE: Reaction Data and substance data are stored in separate documents and can not be searched together in one query. Reaction data for BEILSTEIN compounds may be displayed immediately with the display codes PRE (preparations) and REA (reactions). A substance answer set retrieved after the search for a chemical name, a compounds with available reaction information by combining with PRE/FA, REA/FA or more generally with RX/FA. The BEILSTEIN Registry Number (BRN) is the link between a BEILSTEIN compound and belonging reactions. For more detailed reaction searches BRNs can be searched as reaction partner BRNs. Reactant BRN (RX.RBRN) or Product BRN (RX.PBRN).<<<

>>> FOR SEARCHING PREPARATIONS SEE HELP PRE <<<

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* FOR PRICE INFORMATION SEE HELP COST

>>> Price change as of January 1st, 2008: Connect Time and Structure
Search fees re-introduced. See NEWS and HELP COST <<<

=> D STAT QUE L17
L1 STR



Structure attributes must be viewed using STN Express query preparation.
L6 39 SEA FILE=REGISTRY SSS FUL L1
L17 1 SEA FILE=REILSTEIN SPE=ON ABR=ON PLI=ON 1.6

=> FILE BABS
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FILE LAST UPDATED: 11 MAY 2009 <20090511/UP>
FILE COVERS 1980 TO DATE.

=> D STAT QUE L19
L19 1 SEA FILE=BARS SPE=ON ABB=ON PLI=ON 5704055/BARSAN

=> DUP REM L34 L14 L19 L17
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FILE 'BEILSTEIN' ENTERED AT 15:07:19 ON 02 OCT 2009
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 PROCESSING COMPLETED FOR L34
 PROCESSING COMPLETED FOR L14
 PROCESSING COMPLETED FOR L19
 PROCESSING COMPLETED FOR L17
 L36 17 DUP REM L34 L14 L19 L17 (0 DUPLICATES REMOVED)
 ANSWERS '1-14' FROM FILE HCAPLUS
 ANSWER '15' FROM FILE WPIX
 ANSWER '16' FROM FILE BABS
 ANSWER '17' FROM FILE BEILSTEIN

=> D IBIB ABS ED HITSTR 1-14; D IBIB AB QHIT 15; D ALL 16; D IDE ALLREF 17

L36 ANSWER 1 OF 17 HCAPLUS COPYRIGHT 2009 ACS on STN
 ACCESSION NUMBER: 2009:703033 HCAPLUS Full-text
 DOCUMENT NUMBER: 151:56726
 TITLE: Nitrogen-containing heterocyclic compounds as
 tachykinin receptor antagonists and their preparation
 and use in the treatment of diseases
 INVENTOR(S): Ikeura, Yoshinori; Shirai, Junya; Sugiyama, Hideyuki;
 Nishikimi, Yuji; Kamei, Taku; Sakauchi, Nobuki
 PATENT ASSIGNEE(S): Takeda Pharmaceutical Company Limited, Japan
 SOURCE: PCT Int. Appl., 471pp.
 CODEN: PIXXD2
 DOCUMENT TYPE: Patent
 LANGUAGE: English
 FAMILY ACC. NUM. COUNT: 1
 PATENT INFORMATION:

| PATENT NO. | KIND | DATE | APPLICATION NO. | DATE |
|---|------|----------|-----------------|----------|
| WO 2009072643 | A1 | 20090611 | WO 2008-JP72224 | 20081202 |
| W: AE, AG, AL, AM, AO, AT, AU, AZ, BA, BB, BG, BH, BR, BW, BY, BZ, CA, CH, CN, CO, CR, CU, CZ, DE, DK, DM, DO, DZ, EC, EE, EG, ES, FI, GB, GD, GE, GH, GM, GT, HN, HR, HU, ID, IL, IN, IS, JP, KE, KG, KM, KN, KP, KR, KZ, LA, LC, LK, LR, LS, LT, LU, LY, MA, MD, ME, MG, MK, MN, MW, MX, MY, MZ, NA, NG, NI, NO, NZ, OM, PG, PH, PL, PT, RO, RS, RU, SC, SD, SE, SG, SK, SL, SM, ST, SV, SY, TJ, TM, TN, TR, TT, TZ, UA, UG, US, UZ, VC, VN, ZA, ZM, ZW | | | | |
| RW: AT, BE, BG, CH, CY, CZ, DE, DK, EE, ES, FI, FR, GB, GR, HR, HU, IE, IS, IT, LT, LU, LV, MC, MT, NL, NO, PL, PT, RO, SE, SI, SK, TR, BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN, TD, TG, BW, GH, GM, KE, LS, MW, MZ, NA, SD, SL, SZ, TZ, UG, ZM, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM | | | | |
| US 20090156572 | A1 | 20090618 | US 2008-314015 | 20081202 |

PRIORITY APPLN. INFO.:

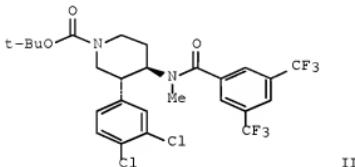
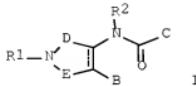
US 2007-996734P

P 20071203

OTHER SOURCE(S):

MARPAT 151:56726

GI



AB The invention relates to a compound represented by formula I, which has a superior tachykinin receptor antagonistic action, and is useful as an agent for the prophylaxis or treatment of various diseases such as lower urinary tract diseases, gastrointestinal diseases, central nervous system diseases and the like. Compds. of formula I wherein B is (un)substituted aromatic ring; C is (un)substituted cyclic group; D is $(CH_2)_n$; and E is $(CH_2)_m$; m and n are independently 0 to 5, and m + n is an integer of 2 to 5; dashed bond is single or double bond; and salts thereof, are claimed. Example compound II was prepared by methylation of tert-Bu $(3R^*,4R^*)$ -4-((1,3-*bis*(trifluoromethyl)phenyl)carbonyl)amino)-3-(3,4- dichlorophenyl)piperidine-1-carboxylate with Me iodide. All the invention compds. were evaluated for their tachykinin receptor antagonistic activity (some data given).

ED Entered STN: 11 Jun 2009

IT 1160255-89-9P

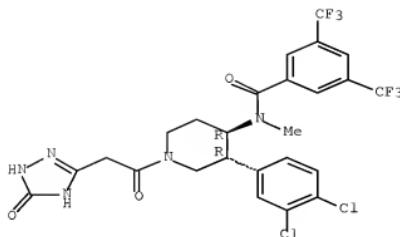
RL: PAC (Pharmacological activity); SPN (Synthetic preparation); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); USES (Uses)
(preparation of nitrogen-containing heterocyclic compds. as tachykinin receptor

antagonists useful in the treatment of diseases)

RN 1160255-89-9 HCPLUS

CN Benzamide, N-((3R,4R)-3-(3,4-dichlorophenyl)-1-[2-(2,5-dihydro-5-oxo-1H-1,2,4-triazol-3-yl)acetyl]-4-piperidinyl)-N-methyl-3,5-*bis*(trifluoromethyl)- (CA INDEX NAME)

Absolute stereochemistry.

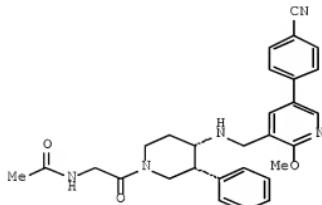
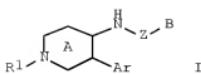


REFERENCE COUNT: 4 THERE ARE 4 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L36 ANSWER 2 OF 17 HCPLUS COPYRIGHT 2009 ACS on STN
 ACCESSION NUMBER: 2007:874350 HCPLUS Full-text
 DOCUMENT NUMBER: 147:257652
 TITLE: Preparation of piperidine derivatives as tachykinin receptor antagonists
 INVENTOR(S): Shirai, Junya; Yoshikawa, Takeshi; Sugiyama, Hideyuki
 PATENT ASSIGNEE(S): Takeda Pharmaceutical Company Limited, Japan
 SOURCE: PCT Int. Appl., 133pp.
 CODEN: PIXXD2
 DOCUMENT TYPE: Patent
 LANGUAGE: English
 FAMILY ACC. NUM. COUNT: 1
 PATENT INFORMATION:

| PATENT NO. | KIND | DATE | APPLICATION NO. | DATE |
|---|------|----------|-----------------|----------|
| WO 2007089031 | A1 | 20070809 | WO 2007-JP52160 | 20070201 |
| W: AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BW, BY, BZ, CA, CH, CN, CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, EG, ES, FI, GB, GD, GE, GH, GM, GT, HN, HR, HU, ID, IL, IN, IS, JP, KE, KG, KM, KN, KP, KR, KZ, LA, LC, LK, LR, LS, LT, LU, LV, LY, MA, MD, MG, MK, MN, MW, MX, MY, MZ, NA, NG, NI, NO, NZ, OM, PG, PH, PL, PT, RO, RS, RU, SC, SD, SE, SG, SK, SL, SM, SV, SY, TJ, TM, TN, TR, TT, TZ, UA, UG, US, UZ, VC, VN, ZA, ZM, ZW | | | | |
| RW: AT, BE, BG, CH, CY, CZ, DE, DK, EE, ES, FI, FR, GB, GR, HU, IE, IS, IT, LT, LU, LV, MC, NL, PL, PT, RO, SE, SI, SK, TR, BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN, TD, TG, BW, GH, GM, KE, LS, MW, MZ, NA, SD, SL, SZ, TZ, UG, ZM, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM | | | | |

PRIORITY APPLN. INFO.: US 2006-763894P P 20060201
 OTHER SOURCE(S): CASREACT 147:257652; MARPAT 147:257652
 GI



AB Title compds. I [Ar = (un)substituted phenyl; R1 = H, (un)substituted hydrocarbyl, acyl or heterocyclyl; Z = (un)substituted methylene; ring A = (un)substituted piperidine; B = (un)substituted monocyclic aromatic heterocyclyl with provisions that substituents may form a ring], and their pharmaceutically acceptable salts, prodrugs are prepared and disclosed as tachykinin receptor antagonists and useful as an agent for the prophylaxis or treatment of lower urinary tract disease and the like. Thus, e.g., II was prepared by condensation of N-[2-((3R,4S)-4-amino-3-phenylpiperidin-1-yl)-2-oxoethyl]acetamide methanesulfonate (preparation given) with 4-(5-formyl-6-methoxy-3-pyridinyl-3-yl)benzonitrile (preparation given) followed by reduction I have superior antagonistic activity, e.g., II showed IC50 value of 0.015 nM.

ED Entered STN: 10 Aug 2007

IT 945954-65-4P 945954-79-0P

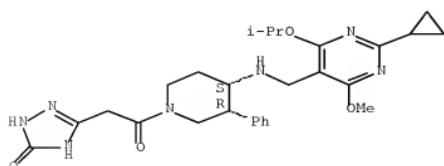
RL: PAC (Pharmacological activity); SPN (Synthetic preparation); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); USES (Uses)

(drug candidate; preparation of piperidine derivs. as tachykinin receptor antagonists)

RN 945954-65-4 HCPLUS

CN 3H-1,2,4-Triazol-3-one, 5-[2-[(3R,4S)-4-[[[2-cyclopropyl-4-methoxy-6-(1-methylethoxy)-5-pyrimidinyl]methyl]amino]-3-phenyl-1-piperidinyl]-2-oxoethyl]-1,2-dihydro- (CA INDEX NAME)

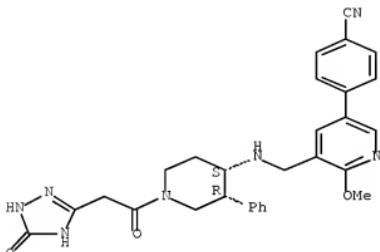
Absolute stereochemistry.



RN 945954-79-0 HCPLUS

CN Benzonitrile, 4-[5-[(3R,4S)-1-[2-(2,5-dihydro-5-oxo-1H-1,2,4-triazol-3-yl)acetyl]-3-phenyl-4-piperidinyl]amino]methyl]-6-methoxy-3-pyridinyl-
(CA INDEX NAME)

Absolute stereochemistry.



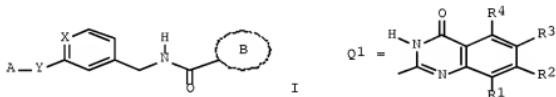
REFERENCE COUNT: 3 THERE ARE 3 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L36 ANSWER 3 OF 17 HCPLUS COPYRIGHT 2009 ACS on STN
 ACCESSION NUMBER: 2007:485967 HCPLUS [Full-text](#)
 DOCUMENT NUMBER: 146:482087
 TITLE: Preparation of heterocyclic amide compounds as matrix metalloproteinase inhibitors
 INVENTOR(S): Nara, Hiroshi; Kaieda, Akira; Sato, Kenjiro; Terauchi, Jun
 PATENT ASSIGNEE(S): Takeda Pharmaceutical Company Limited, Japan
 SOURCE: PCT Int. Appl., 330pp.
 CODEN: PIXXD2
 DOCUMENT TYPE: Patent
 LANGUAGE: Japanese
 FAMILY ACC. NUM. COUNT: 1
 PATENT INFORMATION:

| PATENT NO. | KIND | DATE | APPLICATION NO. | DATE |
|---|------|----------|------------------|----------|
| WO 2007049820 | A1 | 20070503 | WO 2006-JP322043 | 20061027 |
| W: AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BW, BY, BZ, CA, CH, CN, CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, EG, ES, FI, GB, GD, GE, GH, GM, GT, HN, HR, HU, ID, IL, IN, IS, JP, KE, KG, KM, KN, KP, KR, KZ, LA, LC, LK, LR, LS, LT, LU, LV, LY, MA, MD, MG, MK, MN, MW, MX, MY, MZ, NA, NG, NI, NO, NZ, OM, PG, PH, PL, PT, RO, RS, RU, SC, SD, SE, SG, SK, SL, SM, SV, SY, TJ, TM, TN, TR, TT, TZ, UA, UG, US, UZ, VC, VN, ZA, ZM, ZW | | | | |
| RW: AT, BE, BG, CH, CY, CZ, DE, DK, EE, ES, FI, FR, GB, GR, HU, IE, IS, IT, LT, LU, LV, MC, NL, PL, PT, RO, SE, SI, SK, TR, BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN, TD, TG, BW, GH, GM, KE, LS, MW, MZ, NA, SD, SL, SZ, TZ, UG, ZM, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM | | | | |

| | | | | |
|---|----|----------|------------------|------------|
| AU 2006306991 | A1 | 20070503 | AU 2006-306991 | 20061027 |
| CA 2627497 | A1 | 20070503 | CA 2006-2627497 | 20061027 |
| EP 1953148 | A1 | 20080806 | EP 2006-822961 | 20061027 |
| R: AT, BE, BG, CH, CY, CZ, DE, DK, EE, ES, FI, FR, GB, GR, HU, IE, IS, IT, LI, LT, LU, LV, MC, NL, PL, PT, RO, SE, SI, SK, TR, AL, BA, HR, MK, RS | | | | |
| MX 2008005416 | A | 20080512 | MX 2008-5416 | 20080425 |
| US 20090137603 | A1 | 20090528 | US 2008-91773 | 20080428 |
| IN 2008KN01865 | A | 20090109 | IN 2008-KN1865 | 20080508 |
| KR 2008066061 | A | 20080715 | KR 2008-712886 | 20080528 |
| NO 2008002411 | A | 20080728 | NO 2008-2411 | 20080528 |
| CN 101351453 | A | 20090121 | CN 2006-80049861 | 20080630 |
| RITY APPLN. INFO.: | | | | |
| JP 2005-315267 | | | | A 20051028 |
| WO 2006-JP322043 | | | | W 20061027 |

OTHER SOURCE(S): MARPAT 146:482087
GI



AB The title compds. I [A = zinc-bonding group; X = CZ, N; Z = H, halo; Y = (un)substituted spacer having 2 to 10 atoms; ring B = Q1, etc.; R1 - R4 = H, halo, cyano, etc.; excluding 6 specific compds.] are prepared. Thus, 4-oxo-N-[3-((2-((1H-1,2,4-triazol-3-ylthio)ethyl)oxy)phenyl)methyl]-3,4-dihydroquinazoline-2-carboxamide was prepared in several steps starting from 3-hydroxybenzonitrile and 1-bromo-2-chloroethane. In an in vitro assay, compds. of this invention at 1 μ M gave 81% to 100% inhibition of matrix metalloproteinase 13. Formulations are given.

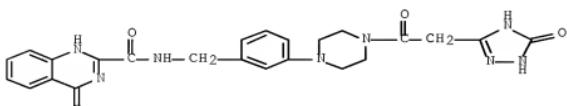
ED Entered STN: 04 May 2007
IT 935759-87-8p

11 553155-8-68
RL: PAC (Pharmacological activity); SPN (Synthetic preparation); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); USES (Uses).

(preparation of heterocyclic amide compds. as matrix metalloproteinase inhibitors)

BN 935759-87-8 HCAPL115

AN 93-39-8
CN 2-Quinazolininecarboxamide, N-[(3-[4-[2-(2,5-dihydro-5-oxo-1H-1,2,4-triazol-3-yl)acetyl]-1-piperazinyl]phenyl)methyl]-3,4-dihydro-4-oxo- (CA INDEX NAME)

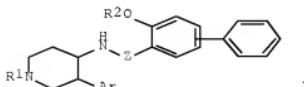


OS.CITING REF COUNT: 1 THERE ARE 1 CAPLUS RECORDS THAT CITE THIS RECORD
(1 CITINGS)
REFERENCE COUNT: 5 THERE ARE 5 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L36 ANSWER 4 OF 17 HCPLUS COPYRIGHT 2009 ACS on STN
ACCESSION NUMBER: 2007:150254 HCPLUS Full-text
DOCUMENT NUMBER: 146:206214
TITLE: Preparation of biphenylmethylaminopiperidines as tachykinin receptor antagonists.
INVENTOR(S): Ikeura, Yoshinori; Shirai, Junya; Yoshikawa, Takeshi; Sakauchi, Nobuki
PATENT ASSIGNEE(S): Takeda Pharmaceutical Company Limited, Japan
SOURCE: PCT Int. Appl., 174pp.
DOCUMENT TYPE: Patent
LANGUAGE: English
FAMILY ACC. NUM. COUNT: 2
PATENT INFORMATION:

| PATENT NO. | KIND | DATE | APPLICATION NO. | DATE |
|---|------|----------|------------------|------------|
| WO 2007015588 | A1 | 20070208 | WO 2006-JP315899 | 20060804 |
| W: AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BW, BY, BZ, CA, CH, CN, CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, EG, ES, FI, GB, GD, GE, GH, GM, HN, HR, HU, ID, IL, IN, IS, JE, KE, KG, KM, KN, KP, KR, KZ, LA, LC, LK, LR, LS, LT, LU, LV, LY, MA, MD, MG, MK, MN, MW, MX, MZ, NA, NG, NI, NO, NZ, OM, PG, PH, PL, PT, RO, RS, RU, SC, SD, SE, SG, SK, SL, SM, SY, TJ, TM, TN, TR, TT, TZ, UA, UG, US, UZ, VC, VN, ZA, ZM, ZW | | | | |
| RW: AT, BE, BG, CH, CY, CZ, DE, DK, EE, ES, FI, FR, GB, GR, HU, IE, IS, IT, LT, LU, LV, MC, NL, PL, PT, RO, SE, SI, SK, TR, BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN, TD, TG, BW, GH, GM, KE, LS, MW, MZ, NA, SD, SL, SZ, TZ, UG, ZM, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM | | | | |
| EP 1910292 | A1 | 20080416 | EP 2006-782685 | 20060804 |
| R: AT, BE, BG, CH, CY, CZ, DE, DK, EE, ES, FI, FR, GB, GR, HU, IE, IS, IT, LI, LT, LU, LV, MC, NL, PL, PT, RO, SE, SI, SK, TR | | | | |
| JP 2009502739 | T | 20090129 | JP 2008-505537 | 20060804 |
| US 20070149570 | A1 | 20070628 | US 2007-701380 | 20070202 |
| PRIORITY APPLN. INFO.: | | | JP 2005-227183 | A 20050804 |
| | | | WO 2006-JP315899 | W 20060804 |

OTHER SOURCE(S): CASREACT 146:206214; MARPAT 146:206214
GI



AB Title compds. [I; Ar = (substituted) Ph; R1 = H, (substituted) hydrocarbyl, acyl, heterocyclyl; R2 = H, (substituted) alkyl, cycloalkyl; Z = (alkyl-

substituted) methylene; all rings may be further substituted; with 2 specifically excluded compds.), were prepared. Thus, N-[2-[(3R,4S)-4-[(4'-ethynyl-4-methoxybiphenyl-3-yl)methyl]amino]-3-phenylpiperidin-1-yl]-2-oxoethyl]acetamide (general preparation given) showed radioligand receptor binding inhibitory activity in IM-9 human lymphoblast cells with IC₅₀ = 0.015 nM.

ED Entered STN: 09 Feb 2007

IT 923280-44-8P 923280-84-6P

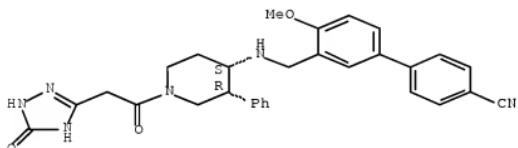
RL: PAC (Pharmacological activity); SPN (Synthetic preparation); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); USES (Uses)

(preparation of biphenylmethylaminopiperidines as tachykinin receptor antagonists)

RN 923280-44-8 HCPLUS

CN [1,1'-Biphenyl]-4-carbonitrile, 3'-{[(3R,4S)-1-[(2-(2,5-dihydro-5-oxo-1H-1,2,4-triazol-3-yl)acetyl)-3-phenyl-4-piperidinyl]amino]methyl}-4'-methoxy- (CA INDEX NAME)

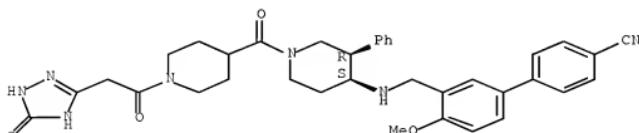
Absolute stereochemistry.



RN 923280-84-6 HCPLUS

CN [1,1'-Biphenyl]-4-carbonitrile, 3'-{[(3R,4S)-1-[(1-[2-(2,5-dihydro-5-oxo-1H-1,2,4-triazol-3-yl)acetyl]-4-piperidinyl)carbonyl]-3-phenyl-4-piperidinyl]amino]methyl}-4'-methoxy-, hydrochloride (1:1) (CA INDEX NAME)

Absolute stereochemistry.



● HCl

REFERENCE COUNT:

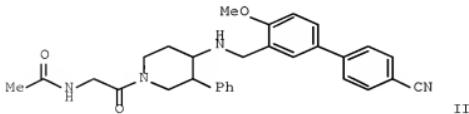
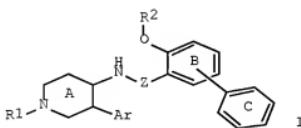
2

THERE ARE 2 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L36 ANSWER 5 OF 17 HCPLUS COPYRIGHT 2009 ACS on STN
ACCESSION NUMBER: 2007:705062 HCPLUS [Full-text](#)

DOCUMENT NUMBER: 147:118148
 TITLE: Piperidine derivatives as tachykinin receptor antagonists and their preparation, pharmaceutical compositions and use in the treatment of lower urinary tract symptoms, gastrointestinal and central nerve disease
 INVENTOR(S): Ikeura, Yoshinori; Shirai, Junya; Yoshikawa, Takeshi; Sakauchi, Nobuki
 PATENT ASSIGNEE(S): Takeda Pharmaceutical Co., Ltd., Japan
 SOURCE: U.S. Pat. Appl. Publ., 89 pp., Cont.-in-part of Appl. No. PCT/JP2006/315899.
 CODEN: USXKCO
 DOCUMENT TYPE: Patent
 LANGUAGE: English
 FAMILY ACC. NUM. COUNT: 2
 PATENT INFORMATION:

| PATENT NO. | KIND | DATE | APPLICATION NO. | DATE |
|---|------|------------------|-----------------|------------|
| US 20070149570 | A1 | 20070628 | US 2007-701380 | 2007070202 |
| WO 2007015588 | A1 | 20070208 | WO 2006-315899 | 20060804 |
| W: AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BW, BY, BZ, CA, CH, CN, CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, EG, ES, FI, GB, GD, GE, GH, GM, HN, HR, HU, ID, IL, IN, IS, JP, KE, KG, KM, KN, KP, KR, KZ, LA, LC, LK, LR, LS, LT, LU, LV, LY, MA, MD, MG, MK, MN, MW, MX, MZ, NA, NG, NI, NO, NZ, OM, PG, PH, PL, PT, RO, RS, RU, SC, SD, SE, SG, SK, SL, SM, SY, TJ, TM, TN, TR, TT, TZ, UA, UG, US, UZ, VC, VN, ZA, ZM, ZW | | | | |
| RW: AT, BE, BG, CH, CY, CZ, DE, DK, EE, ES, FI, FR, GB, GR, HU, IE, IS, IT, LT, LU, LV, MC, NL, PL, PT, RO, SE, SI, SK, TR, BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN, TD, TG, BW, GH, GM, KE, LS, MW, MZ, NA, SD, SL, SZ, TZ, UG, ZM, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM | | | | |
| PRIORITY APPLN. INFO.: | | JP 2005-227183 | A 20050804 | |
| | | WO 2006-JP315899 | A2 20060804 | |
| OTHER SOURCE(S): MARPAT 147:118148 GI | | | | |



AB The invention relates to a compound represented by formula I or a salt thereof. Compds. of formula I wherein Ar is (un)substituted Ph; R1 is H, (un)substituted hydrocarbon, acyl and (un)substituted heterocyclic group; R2 is H, (un)substituted C1-6 alkyl and (un)substituted C3-6 cycloalkyl; Z is (un)substituted methylene; ring A is a (un)substituted piperidine ring; ring B and ring C are (un)substituted benzene; R2 optionally form a ring together with the adjacent substituent on the ring B; and their salts thereof, are claimed. The compound of the invention has a superior tachykinin receptor antagonistic action, particularly a substance P receptor antagonistic action, and is useful as a pharmaceutical agent, for example, tachykinin receptor antagonist, an agent for the prophylaxis or treatment of lower urinary tract symptoms, gastrointestinal diseases or central nerve diseases. Example compound II was prepared by a general procedure (procedure given). All the invention compds. were evaluated for their tachykinin receptor antagonistic activity. From the assay, it was determined that compound II exhibited an IC50 value of 0.019 nM.

ED Entered STN: 29 Jun 2007

IT 923280-44-8P 923280-84-6P

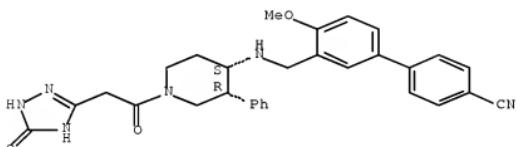
RL: PAC (Pharmacological activity); SPN (Synthetic preparation); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); USES (Uses)

(drug candidate; preparation of piperidine derivs. as tachykinin receptor antagonists and their use in the treatment of lower urinary tract symptoms, gastrointestinal and central nerve disease)

RN 923280-44-8 HCPLUS

CN [1,1'-Biphenyl]-4-carbonitrile, 3'-([(3R,4S)-1-[(2-(2,5-dihydro-5-oxo-1H-1,2,4-triazol-3-yl)acetyl]-3-phenyl-4-piperidinyl]amino]methyl)-4'-methoxy- (CA INDEX NAME)

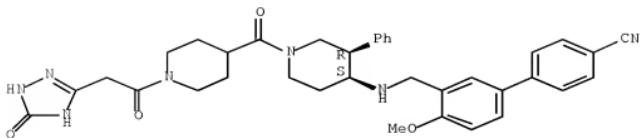
Absolute stereochemistry.



RN 923280-84-6 HCPLUS

CN [1,1'-Biphenyl]-4-carbonitrile, 3'-([(3R,4S)-1-[(1-[2-(2,5-dihydro-5-oxo-1H-1,2,4-triazol-3-yl)acetyl]-4-piperidinyl]carbonyl]-3-phenyl-4-piperidinyl]amino)methyl]-4'-methoxy-, hydrochloride (1:1) (CA INDEX NAME)

Absolute stereochemistry.



HCL

L36 ANSWER 6 OF 17 HCPLUS COPYRIGHT 2009 ACS on STN
ACCESSION NUMBER: 2006:1155411 HCPLUS Full-text
DOCUMENT NUMBER: 145:471540
TITLE: Preparation of piperidine derivatives as tachykinin receptor antagonists
INVENTOR(S): Nagaoka, Naomi; Marunaka, Shigeyuki; Fukuta, Makoto
PATENT ASSIGNEE(S): Takeda Pharmaceutical Company Limited, Japan
SOURCE: PCT Int. Appl., 323pp.
CODEN: PIXXD2
DOCUMENT TYPE: Patent
LANGUAGE: Japanese
FAMILY ACC. NUM. COUNT: 1
PATENT INFORMATION:

| PATENT NO. | KIND | DATE | APPLICATION NO. | DATE |
|---|------|----------|------------------|----------|
| WO 2006115285 | A1 | 20061102 | WO 2006-JP308919 | 20060421 |
| W: AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BW, BY, BZ, CA, CH, CN, CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, EG, ES, FI, GB, GD, GE, GH, GM, HR, IU, ID, IL, IN, IS, JP, KE, KG, KM, KN, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, LY, MA, MD, MG, MK, MN, MW, MX, MZ, NA, NG, NI, NO, NZ, OM, PG, PH, PL, PT, RO, RU, SC, SD, SE, SG, SK, SL, SM, SY, TJ, TM, TN, TR, TT, TZ, UA, UG, US, UZ, VC, VN, YU, ZA, ZM, ZW | | | | |
| RW: AT, BE, BG, CH, CY, CZ, DE, DK, EE, ES, FI, FR, GB, GR, HU, IE, IS, IT, LT, LU, LV, MC, NL, PL, PT, RO, SE, SI, SK, TR, BF, BJ, CF, CG, CI, CM, GA, GN, GO, GM, ML, MR, NE, SN, SK, TD, TG, BW, GH, GM, KE, LS, MW, MZ, NA, SD, SL, SZ, TZ, UG, ZM, ZW, AM, AZ, BY, KC, KZ, MD, RU, TL, TM | | | | |

PRIO RY APPN INFO : 1P 2005-124335 A 20050421

OTHER SOURCE(S): MARRAT 145:471540

AB The title compds. (no biol. data) are prepared. This document discloses a pharmaceutical composition comprising N-(2-[(3R,4S)-4-((2-methoxy-5-[trifluoromethyl]-1H-tetrazol-1-yl)benzyl)amino]-3-phenylpiperidin-1-yl]-2-oxoethyl)acetamide (I), a salt or a prodrug thereof, a sugar and a hydrophilic water-insol. substance. Thus, N-(2-[(3R,4S)-4-((2-hydroxy-5-[trifluoromethyl]-1H-tetrazol-1-yl)benzyl)amino]-3-phenylpiperidin-1-yl]-2-oxoethyl)acetamide was prepared in 3 steps from (3R,4S)-4-amino-3-phenylpiperidine-1-carboxylic acid tert-Bu ester and 2-hydroxy-5-[trifluoromethyl]-1H-tetrazol-1-yl)benzaldehyde. Formulations containing I are given. Tablets containing I showed high elution stability.

ED Entered STN: 03 Nov 2006

IT 632352-46-6P

RL: PAC (Pharmacological activity); SPN (Synthetic preparation); THU

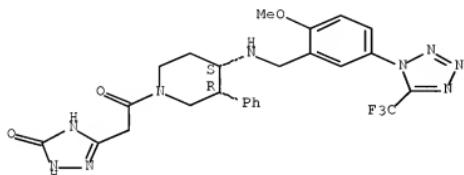
(Therapeutic use); BIOL (Biological study); PREP (Preparation); USES (Uses)

(preparation of piperidine derivs. as tachykinin receptor antagonists)

RN 632352-46-6 HCPLUS

CN 3H-1,2,4-Triazol-3-one, 1,2-dihydro-5-[2-[(3R,4S)-4-[(2-methoxy-5-[5-(trifluoromethyl)-1H-tetrazol-1-yl]phenyl)methyl]amino]-3-phenyl-1-piperidinyl]-2-oxoethyl]-, rel- (CA INDEX NAME)

Relative stereochemistry.

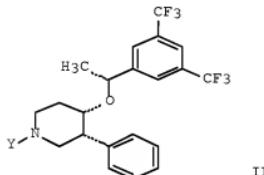
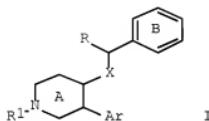


REFERENCE COUNT: 36 THERE ARE 36 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L36 ANSWER 7 OF 17 HCPLUS COYRIGHT 2009 ACS on STN
 ACCESSION NUMBER: 2006:272922 HCPLUS Full-text
 DOCUMENT NUMBER: 144:331270
 TITLE: Preparation of piperidine derivatives as tachykinin receptor antagonists
 INVENTOR(S): Ikeura, Yoshinori; Hashimoto, Tadatoshi; Nishida, Haruyuki; Shirai, Junya; Sakauchi, Nobuki
 PATENT ASSIGNEE(S): Takeda Pharmaceutical Company Limited, Japan
 SOURCE: PCT Int. Appl., 222 pp.
 CODEN: PIXXD2
 DOCUMENT TYPE: Patent
 LANGUAGE: Japanese
 FAMILY ACC. NUM. COUNT: 1
 PATENT INFORMATION:

| PATENT NO. | KIND | DATE | APPLICATION NO. | DATE |
|---|------|----------|-----------------|----------|
| WO 2006030975 | A1 | 20060323 | WO 2005-JP17538 | 20050916 |
| W: AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BW, BY, BZ, CA, CH, CN, CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, EG, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KM, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, LY, MA, MD, MG, MK, MN, MW, MX, MZ, NA, NG, NI, NO, NZ, OM, PG, PH, PL, PT, RO, RU, SC, SD, SE, SG, SK, SL, SM, SY, TJ, TM, TN, TR, TT, TZ, UA, UG, US, UZ, VC, VN, YU, ZA, ZM, ZW | | | | |
| RW: AT, BE, BG, CH, CY, CZ, DE, DK, EE, ES, FI, FR, GB, GR, HU, IE, IS, IT, LT, LU, LV, MC, NL, PL, PT, RO, SE, SI, SK, TR, BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN, TD, TG, BM, GH, GM, KE, LS, MW, MZ, NA, SD, SL, SZ, TZ, UG, ZM, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM | | | | |
| EP 1790636 | A1 | 20070530 | EP 2005-785870 | 20050916 |
| R: AT, BE, BG, CH, CY, CZ, DE, DK, EE, ES, FI, FR, GB, GR, HU, IE, IS, IT, LI, LT, LU, LV, MC, NL, PL, PT, RO, SE, SI, SK, TR | | | | |

| | | | | |
|------------------------|----|-------------------|-----------------|------------|
| US 20060142337 | A1 | 20060629 | US 2006-358070 | 20060222 |
| PRIORITY APPLN. INFO.: | | | JP 2004-272639 | A 20040917 |
| OTHER SOURCE(S): | | MARPAT 144:331270 | WO 2005-JP17538 | W 20050916 |
| GI | | | | |



AB Title compds. I [Ar = (un)substituted aryl; R = alkyl; R1 = H, (un)substituted hydrocarbon, acyl, etc.; X = O, (un)substituted imino; ring A = piperidine ring which may have an addnl. substituent; ring B = substituted benzene] were prepared. For example, compound II [Y = H]·HCl was prepared from (3R,4S)-4-hydroxy-3-phenylpiperidine-1-carboxylic acid tert-Bu ester in a multistep process. In radioligand receptor binding inhibition assays, compound II [Y = (1-acetylpiriperidin-4-yl)carbonyl] exhibited the IC50 value of 0.026 nM. Compds. I are claimed useful for the treatment of irritable bowel disease, depression, etc.

ED Entered STN: 24 Mar 2006

IT 880092-22-8P 880092-48-8P 880092-89-7P

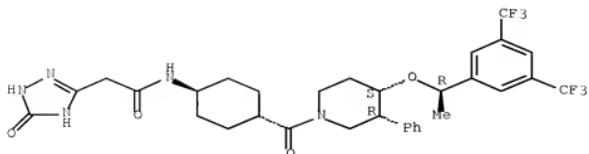
RL: PAC (Pharmacological activity); SPN (Synthetic preparation); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); USES (Uses)

(preparation of piperidine derivs. as tachykinin receptor antagonists for treatment of irritable bowel disease, depression, etc.)

RN 880092-22-8 HCPLUS

CN 1H-1,2,4-Triazole-3-acetamide, N-[trans-4-[(3R,4S)-4-[(1R)-1-[3,5-bis(trifluoromethyl)phenyl]ethoxy]-3-phenyl-1-piperidinyl]carbonyl]cyclohexyl]-2,5-dihydro-5-oxo- (CA INDEX NAME)

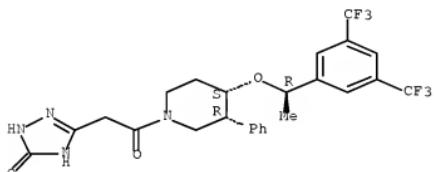
Absolute stereochemistry.



RN 880092-48-8 HCPLUS

CN 3H-1,2,4-Triazol-3-one, 5-[2-[(3R,4S)-4-[(1R)-1-[3,5-bis(trifluoromethyl)phenyl]ethoxy]-3-phenyl-1-piperidinyl]-2-oxoethyl]-1,2-dihydro- (CA INDEX NAME)

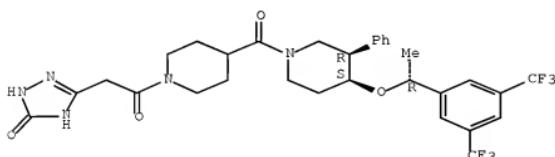
Absolute stereochemistry.



RN 880092-89-7 HCPLUS

CN 3H-1,2,4-Triazol-3-one, 5-[2-[(3R,4S)-4-[(1R)-1-[3,5-bis(trifluoromethyl)phenyl]ethoxy]-3-phenyl-1-piperidinyl]carbonyl]-1-piperidinyl]-2-oxoethyl]-1,2-dihydro- (CA INDEX NAME)

Absolute stereochemistry.

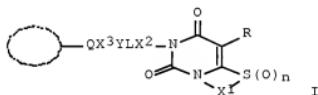
OS.CITING REF COUNT: 3 THERE ARE 3 CAPLUS RECORDS THAT CITE THIS RECORD
(3 CITINGS)

REFERENCE COUNT: 12 THERE ARE 12 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L36 ANSWER 8 OF 17 HCPLUS COPYRIGHT 2009 ACS on STN
ACCESSION NUMBER: 2004:472319 HCPLUS [Full-text](#)

DOCUMENT NUMBER: 141:47322
TITLE: Sulfur heterocycle-condensed pyrimidinedione derivatives, prodrugs of them, JNK inhibitors containing them, and pharmaceuticals containing them
INVENTOR(S): Ito, Fumio; Kimura, Hiroyuki; Ikata, Hideki; Kitamura, Shuji; Kawamoto, Tomohiro; Abe, Hidenori
PATENT ASSIGNEE(S): Takeda Chemical Industries, Ltd., Japan
SOURCE: Jpn. Kokai Tokyo Koho, 117 pp.
CODEN: JKXXAF
DOCUMENT TYPE: Patent
LANGUAGE: Japanese
FAMILY ACC. NUM. COUNT: 1
PATENT INFORMATION:

| PATENT NO. | KIND | DATE | APPLICATION NO. | DATE |
|------------------------|--------|-----------|-----------------|----------|
| JP 2004161716 | A | 20040610 | JP 2002-332027 | 20021115 |
| PRIORITY APPLN. INFO.: | | | JP 2002-332027 | 20021115 |
| OTHER SOURCE(S): | MARPAT | 141:47322 | | |
| CT | | | | |



AB The derivs., useful for prevention and treatment of diseases involving JNK, e.g. cardiac failure, hypertension, rheumatoid arthritis, asthma, Alzheimer's disease, ischemia, etc., are represented by I [$R = H$, (un)substituted hydrocarbyl, (un)substituted heterocycl; $X_1, X_2 =$ (un)substituted C2-4 alkylene; $X_3 =$ direct bond, (un)substituted C1-5 alkylene, (un)substituted C2-4 alkylene; $Y =$ direct bond, (un)substituted divalent cyclic group; $Q =$ direct bond, O, S, NR1 [$R_1 = H$, (un)substituted lower alkyl]; L = direct bond, CONR2 [$R_2 = H$, (un)substituted lower alkyl]; ring A = (un)substituted N-heterocycle; $n = 0, 1, 2$. JNK inhibitors contain I, their salts, or prodrugs of I. Thus, IC50 of 4-(6-aminopyridin-3-yl)-N-[3-(1,1,6,8-tetraoxo-9-phenyl-1,3,4,8-tetrahydro-2H-1*A*6-pyrimido[6,1-*b*][1,3]thiazin-7-yl)propyl]benzamide hydrochloride (II preparation given) against human JNK1 was 0.00082 μ M. Capsules and tablets containing II were also formulated.

ED Entered STN: 11 Jun 2004

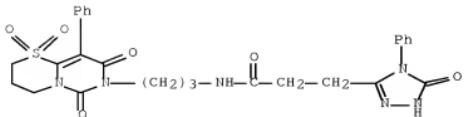
701215-97-6B

RL: PAC (Pharmacological activity); SPN (Synthetic preparation); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); USES (Uses)

(preparation of sulfur heterocycle-condensed pyrimidinedione derivs. as JNK inhibitors)

RN 701215-97-6 HCAPLUS

CN 1H-1,2,4-Triazole-3-propanamide, N-[3-(3,4-dihydro-1,1-dioxido-6,8-dioxo-9-phenyl-2H,6H-pyrimido[6,1-b][1,3]thiazin-7(8H)-yl)propyl]-4,5-dihydro-5-oxo-4-phenyl- (CA INDEX NAME)



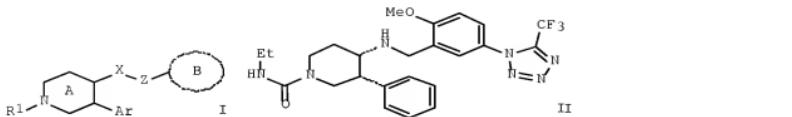
OS.CITING REF COUNT: 3 THERE ARE 3 CAPLUS RECORDS THAT CITE THIS RECORD
(3 CITINGS)

L36 ANSWER 9 OF 17 HCPLUS COPYRIGHT 2009 ACS on STN
ACCESSION NUMBER: 2003:972057 HCPLUS Full-text
DOCUMENT NUMBER: 140:27765
TITLE: Preparation of piperidine derivatives as tachykinin receptor antagonists for treatment of frequent urination and urinary incontinence
INVENTOR(S): Ikeura, Yoshinori; Hashimoto, Tadatoshi; Tarui, Naoki; Shirai, Junya; Yamashita, Masayuki
PATENT ASSIGNEE(S): Takeda Chemical Industries, Ltd., Japan
SOURCE: PCT Int. Appl., 264 pp.
CODEN: PIXXD2
DOCUMENT TYPE: Patent
LANGUAGE: Japanese
FAMILY ACC. NUM. COUNT: 1
PATENT INFORMATION:

| PATENT NO. | KIND | DATE | APPLICATION NO. | DATE |
|---|------|----------|-----------------|------------|
| WO 2003101964 | A1 | 20031211 | WO 2003-JP6754 | 20030529 |
| W: AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, BZ, CA, CH, CN, CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NI, NO, NZ, OM, PH, PL, PT, RO, RU, SC, SD, SE, SG, SK, SL, TJ, TM, TN, TR, TT, TZ, UA, US, UZ, VC, VN, YU, ZA, ZM, ZW | | | | |
| RW: GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZM, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM, AT, BE, BG, CH, CY, CZ, DE, DK, EE, ES, FI, FR, GB, GR, HU, IE, IT, LU, MC, NL, PT, RO, SE, SI, SK, TR, BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN, TD, TG | | | | |
| CA 2487688 | A1 | 20031211 | CA 2003-2487688 | 20030529 |
| AU 2003241903 | A1 | 20031219 | AU 2003-241903 | 20030529 |
| BR 2003011425 | A | 20050315 | BR 2003-11425 | 20030529 |
| EP 1553084 | A1 | 20050713 | EP 2003-733151 | 20030529 |
| R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT, IE, SI, LT, LV, FI, RO, MK, CY, AL, TR, BG, CZ, EE, HU, SK | | | | |
| CN 1671662 | A | 20050921 | CN 2003-818354 | 20030529 |
| NZ 537330 | A | 20070427 | NZ 2003-537330 | 20030529 |
| JP 2004285038 | A | 20041014 | JP 2003-154345 | 20030530 |
| MX 2004011730 | A | 20050714 | MX 2004-11730 | 20041125 |
| US 20060167052 | A1 | 20060727 | US 2004-516252 | 20041129 |
| ZA 2004010085 | A | 20060726 | ZA 2004-10085 | 20041214 |
| IN 2004KN01942 | A | 20061201 | IN 2004-KN1942 | 20041216 |
| NO 2004005701 | A | 20050216 | NO 2004-5701 | 20041229 |
| PRIORITY APPLN. INFO.: | | | | |
| | | | JP 2002-159338 | A 20020531 |
| | | | JP 2003-17885 | A 20030127 |
| | | | WO 2003-JP6754 | W 20030529 |

OTHER SOURCE(S) :
GI

MARPAT 140:27765



AB The title compds. I [wherein Ar = (un)substituted aryl, aralkyl, or heteroaryl; R1 = H, acyl, (un)substituted hydrocarbyl, or heterocycl; X = O or (un)substituted NH; Z = (un)substituted CH2; ring A = (un)substituted piperidine; ring B = (un)substituted aryl; with exclusions] or prodrugs or salts thereof are prepared. I have excellent tachykinin receptor antagonistic activity, and are useful for the treatment of frequent urination and urinary incontinence (no data). For example, the compound II•xHCl was prepared in a multi-step synthesis. II showed antagonistic activity with IC50 of 0.025 nM against human substance P receptor. Formulations containing I as an active ingredient were also described.

ED Entered STN: 14 Dec 2003

IT 632352-46-6P

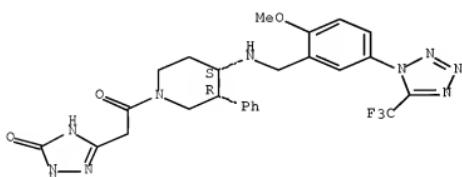
RL: PAC (Pharmacological activity); SPN (Synthetic preparation); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); USES (Uses)

(drug candidate; preparation of piperidine derivs. as tachykinin receptor antagonists for treatment of frequent urination and urinary incontinence)

RN 632352-46-6 HCAPLUS

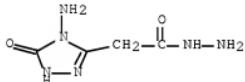
CN 3H-1,2,4-Triazol-3-one, 1,2-dihydro-5-[2-[(3R,4S)-4-[[[2-methoxy-5-[5-(trifluoromethyl)-1H-tetrazol-1-yl]phenyl]methyl]amino]-3-phenyl-1-piperidinyl]-2-oxoethyl]-, rel- (CA INDEX NAME)

Relative stereochemistry.



OS.CITING REF COUNT: 12 THERE ARE 12 CAPLUS RECORDS THAT CITE THIS RECORD (31 CITINGS)
 REFERENCE COUNT: 14 THERE ARE 14 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

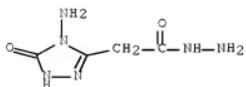
ACCESSION NUMBER: 1997:281511 HCAPLUS Full-text
 DOCUMENT NUMBER: 127:17627
 ORIGINAL REFERENCE NO.: 127:3561a,3564a
 TITLE: Synthesis and antibacterial activities of some
 4,5-dihydro-1H-1,2,4-triazol-5-ones
 AUTHOR(S): Yuksek, Haydar; Demirbas, Ahmet; Ikizler, Aysun;
 Johansson, Candan Bozok; Celik, Cennet; Ikizler, Aykut
 A.
 CORPORATE SOURCE: Department Chemistry, Karadeniz Technical University,
 Trabzon, 61080, Turk.
 SOURCE: Arzneimittel-Forschung (1997), 47(4), 405-409
 CODEN: ARZNAD; ISSN: 0004-4172
 PUBLISHER: Cantor
 DOCUMENT TYPE: Journal
 LANGUAGE: English
 AB A series of 3,4-disubstituted 1-methyl- and 1-ethyl-4,5-dihydro-1H-1,2,4-triazol-5-ones was prepared by reaction of the appropriate dihydrotriazolones with Me2SO4 or Et2SO4. Some of these new and some recently reported 4,5-dihydro-1H-1,2,4-triazol-5-ones exhibited antibacterial and tuberculostatic activities.
 ED Entered STN: 02 May 1997
 IT 75989-59-2
 RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); BIOL (Biological study)
 (antibacterial activity)
 RN 75989-59-2 HCAPLUS
 CN 1H-1,2,4-Triazole-3-acetic acid, 4-amino-4,5-dihydro-5-oxo-, hydrazide
 (CA INDEX NAME)



OS.CITING REF COUNT: 64 THERE ARE 64 CAPLUS RECORDS THAT CITE THIS RECORD (64 CITINGS)

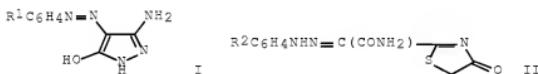
L36 ANSWER 11 OF 17 HCAPLUS COPYRIGHT 2009 ACS on STN
 ACCESSION NUMBER: 1992:173458 HCAPLUS Full-text
 DOCUMENT NUMBER: 116:173458
 ORIGINAL REFERENCE NO.: 116:29339a,29342a
 TITLE: pK'a values of some 1,2,4-triazole derivatives in
 nonaqueous media
 AUTHOR(S): Ikizler, A. Aykut; Senturk, H. Basri; Ikizler, Aysun
 CORPORATE SOURCE: Dep. Chem., Karadeniz Tech. Univ., Trabzon, Turk.
 SOURCE: Doga: Turk Kimya Dergisi (1991), 15(4), 345-54
 CODEN: DKSEE7; ISSN: 1010-7614
 DOCUMENT TYPE: Journal
 LANGUAGE: English
 AB The pK'a values of some 1,2,4-triazole derivs. were determined
 potentiometrically by using 2-propanol solvent and tetrabutylammonium
 hydroxide (TBAH) in 2-propanol as titrant.
 ED Entered STN: 03 May 1992
 IT 75989-59-2
 RL: PRP (Properties)

(acidity of, in isopropanol)
 RN 75989-59-2 HCPLUS
 CN 1H-1,2,4-Triazole-3-acetic acid, 4-amino-4,5-dihydro-5-oxo-, hydrazide
 (CA INDEX NAME)

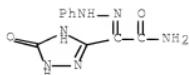


OS.CITING REF COUNT: 14 THERE ARE 14 CAPLUS RECORDS THAT CITE THIS RECORD (14 CITINGS)

L36 ANSWER 12 OF 17 HCPLUS COPYRIGHT 2009 ACS on STN
 ACCESSION NUMBER: 1988:492870 HCPLUS Full-text
 DOCUMENT NUMBER: 109:92870
 ORIGINAL REFERENCE NO.: 109:15497a,15500a
 TITLE: Synthesis of azoles and fused azoles from α -arylhydrazononitriles
 AUTHOR(S): Ibrahim, Mohamed Kamal Ahmed; El-Moghayar, Mohamed Riffat Hamza
 CORPORATE SOURCE: Fac. Sci., Cairo Univ., Giza, Egypt
 SOURCE: Indian Journal of Chemistry, Section B: Organic Chemistry Including Medicinal Chemistry (1987), 26B(9), 832-5
 DOCUMENT TYPE: CODEN: IJSBDB; ISSN: 0376-4699
 LANGUAGE: Journal
 OTHER SOURCE(S): English
 GI: CASREACT 109:92870

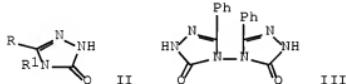


AB Cyanoacetamides $R1C6H4NHN:C(C(=O)CN$ ($R1 = H, Me, Cl$) were heated with $N2H4$ to give pyrazoles I. Also prepared, from cyanoacetamides and $HSCH2CO2H$, were thiazolinones II ($R2 = Cl, CO2H$).
 ED Entered STN: 17 Sep 1988
 IT 115998-45-3P
 RL: SPN (Synthetic preparation); PREP (Preparation)
 (preparation of)
 RN 115998-45-3 HCPLUS
 CN 1H-1,2,4-Triazole-3-acetamide, 2,5-dihydro-5-oxo- α -(2-phenylhydrazinylidene)- (CA INDEX NAME)



OS.CITING REF COUNT: 1 THERE ARE 1 CAPLUS RECORDS THAT CITE THIS RECORD
(2 CITINGS)

L36 ANSWER 13 OF 17 HCPLUS COPYRIGHT 2009 ACS on STN
ACCESSION NUMBER: 1981:15645 HCPLUS Full-text
DOCUMENT NUMBER: 94:15645
ORIGINAL REFERENCE NO.: 94:2619a,2622a
TITLE: Reactions of ester ethoxycarbonylhydrazones with some
amine type compounds
AUTHOR(S): Ikizler, Aykut; Un, Resat
CORPORATE SOURCE: Fac. Chem., Ege Univ., Izmir, Turk.
SOURCE: Chimica Acta Turcica (1979), 7(3), 269-90
CODEN: CATUA9; ISSN: 0379-5896
DOCUMENT TYPE: Journal
LANGUAGE: English
OTHER SOURCE(S): CASREACT 94:15645
GI



AB EtOCR:NNHCO2Et (I, R = Ph, Me, Et, Pr, Me2CH2CH2, PhCH2, p-MeC6H4, CH2CO2Et) were cyclized with H2NNH2·H2O to give the triazolinones II (R1 = NH2), which were condensed with PhCHO to give II' (R1 = N:CHPh). I and PhHNHNH2 similarly gave II (R1 = PhNH). II (R1 = NH2) were converted to several derivs. e.g. II (R1 = NH2Bz) and III. I (R = Me, Ph) reacted with HOCH2CH2NH2 to give II (R1 = CH2CH2OH).

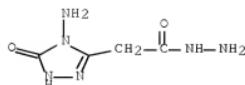
ED Entered STN: 12 May 1984

IT 75989-59-2B

RL: SPN (Synthetic preparation); PREP (Preparation)
(preparation of)

RN 75989-59-2 HCAPLUS

CN 1H-1,2,4-Triazole-3-acetic acid, 4-amino-4,5-dihydro-5-oxo-, hydrazide
(CA INDEX NAME)

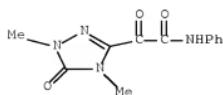


OS.CITING REF COUNT: 39 THERE ARE 39 CAPLUS RECORDS THAT CITE THIS RECORD (43 CITINGS)

L36 ANSWER 14 OF 17 HCAPLUS COPYRIGHT 2009 ACS on STN
 ACCESSION NUMBER: 1977:468245 HCAPLUS Full-text
 DOCUMENT NUMBER: 87:68245
 ORIGINAL REFERENCE NO.: 87:10865a,10868a
 TITLE: Structural elucidation of the reaction products from benzonitrile oxide and 1,4-disubstituted urazoles
 HOYER, Georg A.; BOROSCHEWSKI, Gerhard
 AUTHOR(S):
 CORPORATE SOURCE: Forschungslab., Schering A.-G., Berlin, Fed. Rep. Ger.
 SOURCE: Archiv der Pharmazie (Weinheim, Germany) (1977),
 310(3), 255-9
 DOCUMENT TYPE: CODEN: ARPMA; ISSN: 0365-6233
 LANGUAGE: Journal
 German
 GI



AB The reaction of benzonitrile oxide with urazoles (I; R = R1 = Me; R = Ph, R1 = Me; R = Me, R1 = Ph; R = R1 = Ph) does not yield the corresponding 1,4-disubstituted 3-(phenylcarbamoyloxy)-Δ2-1,2,4-triazolin-5-ones as previously reported (Sunderdiek, R. et al, 1974), but leads to oxadiazolinones (II; R, R1 as above).
 ED Entered STN: 12 May 1984
 IT 63425-53-6
 RL: RCT (Reactant); RACT (Reactant or reagent)
 (oxadiazolinones vs., as reaction products of benzonitrile oxide and urazoles)
 RN 63425-53-6 HCAPLUS
 CN 1H-1,2,4-Triazole-3-acetamide, 4,5-dihydro-1,4-dimethyl- α ,5-dioxo-N-phenyl- (CA INDEX NAME)



L36 ANSWER 15 OF 17 WPIX COPYRIGHT 2009 THOMSON REUTERS on STN
 ACCESSION NUMBER: 2005-734179 [75] WPIX
 DOC. NO. CPI: C2005-223965 [75]
 TITLE: New triazolone derivatives useful for the treatment of
 obstructive airways disease e.g. asthma or chronic
 obstructive pulmonary disease
 DERNWENT CLASS: B02; B03
 INVENTOR: ERIKSSON A; LEPISTOE M; LEPISTO M
 PATENT ASSIGNEE: (ASTR-C) ASTRAZENECA AB; (ERIK-I) ERIKSSON A; (LEPI-I)
 LEPISTO M
 COUNTRY COUNT: 108

PATENT INFO ABBR.:

| PATENT NO | KIND | DATE | WEEK | LA | PG | MAIN IPC |
|-----------------|------|----------|-----------|----|-------|----------|
| WO 2005095362 | A1 | 20051013 | (200575)* | EN | 53[0] | |
| EP 1732903 | A1 | 20061220 | (200702) | EN | | |
| CN 1960979 | A | 20070509 | (200760) | ZH | | |
| US 20070219217 | A1 | 20070920 | (200763) | EN | | |
| IN 2006DN05541 | P1 | 20070803 | (200771) | EN | | |
| JP 2007530672 | W | 20071101 | (200780) | JA | 35 | |
| EP 1732903 | B1 | 20090218 | (200914) | EN | | |
| DE 602005012811 | E | 20090402 | (200927) | DE | | |
| ES 2320679 | T3 | 20090527 | (200943) | ES | | |

APPLICATION DETAILS:

| PATENT NO | KIND | APPLICATION | DATE |
|-----------------------------------|------|----------------------|----------|
| WO 2005095362 A1 | | WO 2005-SE448 | 20050329 |
| IN 2006DN05541 P1 | | WO 2004-SE448 | 20050329 |
| CN 1960979 A | | CN 2005-80017672 | 20050329 |
| DE 602005012811 E | | DE 2005-602005012811 | 20050329 |
| EP 1732903 A1 | | EP 2005-722275 | 20050329 |
| EP 1732903 B1 | | EP 2005-722275 | 20050329 |
| DE 602005012811 E | | EP 2005-722275 | 20050329 |
| EP 1732903 A1 | | WO 2005-SE448 | 20050329 |
| US 20070219217 A1 | | WO 2005-SE448 | 20050329 |
| JP 2007530672 W | | WO 2005-SE448 | 20050329 |
| EP 1732903 B1 PCT Application | | WO 2005-SE448 | 20050329 |
| DE 602005012811 E PCT Application | | WO 2005-SE448 | 20050329 |
| US 20070219217 A1 | | US 2006-593543 | 20060920 |
| IN 2006DN05541 P1 | | IN 2006-DN5541 | 20060922 |
| JP 2007530672 W | | JP 2007-506108 | 20050329 |
| ES 2320679 T3 | | EP 2005-722275 | 20050329 |

FILING DETAILS:

| PATENT NO | KIND | PATENT NO |
|-------------------|----------|-----------------|
| DE 602005012811 E | Based on | EP 1732903 A |
| EP 1732903 | A1 | WO 2005095362 A |
| JP 2007530672 W | Based on | WO 2005095362 A |
| EP 1732903 | B1 | WO 2005095362 A |

| | | |
|-------------------|----------|-----------------|
| DE 602005012811 E | Based on | WO 2005095362 A |
| ES 2320679 T3 | Based on | EP 1732903 A |

PRIORITY APPLN. INFO: SE 2004-850 20040330

AB WO 2005095362 A1 UPAB: 20090307

NOVELTY - Triazolone derivatives are new.

DETAILED DESCRIPTION - Triazolone derivatives of formula (I) or its salt or solvate are new.

R1,R2 = H or 1-6C alkyl (optionally substituted by an aryl ring or an aromatic heterocyclic ring containing 1-3 heteroatoms selected from O, S or N (optionally substituted by halo, CF₃, 1-4C alkyl or 1-4C alkoxy));

R3,R4 = H or 1-6C alkyl (optionally substituted by OH, 1-4C alkoxy, 1-4C alkylthio, amino, N-alkylamino or N,N-dialkylamino);

R3+R4 = 3-7 membered ring (optionally incorporating one heteroatom selected from O, S(O)q or N);

m = 1-3;

X = S(O), S(O)2 or C(=O);

R5 = H or 1-6C alkyl (optionally substituted by halo, OH or 1-6C alkoxy);

Y = direct bond;

NR5 = 4-7 membered optionally saturated azacyclic ring (optionally incorporating one further heteroatom selected from O, S(O)n or N and optionally benzo fused and optionally substituted by 1-6C alkyl, 1-6C alkoxy or OH);

L = direct bond or 2-6C alkynyl, 2-6C alkenyl, 1-6C (hetero)alkyl, or 3-6C heteroalkynyl (all optionally substituted by halo, OH or 1-6C alkoxy), O, S(O)p, C(O), NR6, C(O)NR6 or NR6C(O);

n,p,q = 0-2;

G1 = 1-4 membered monocyclic, bicyclic, tricyclic or tetracyclic group (where each ring structures is of 7 ring atoms) selected from cycloalkyl, cycloalkenyl, optionally saturated heterocycloalkyl (where all alkyl is optionally substituted by halo, OH, 1-6C alkyl, 1-6C alkoxy, halo-(1-6C) alkoxy, amino, N-alkylamino, N,N-dialkylamino, N-alkylsulfonamino, N-2-6C alkanoylamino, cyano, nitro, thiol, alkylthio, alkylsulfonyl, N-alkylaminosulfonyl, CHO, 2-6C alkanoyl, aminocarbonyl, N-alkylaminocarbonyl or N,N-dialkylaminocarbonyl or carbamate), aryl, or aromatic heterocyclic ring containing 1-3 heteroatoms selected from O, S or N (where each ring is optionally substituted by at least one of halo, OH, CHO, 1-6C alkyl, 1-6C alkoxy, halo-(1-6C) alkoxy, amino, N-alkylamino, N,N-dialkylamino, alkylsulfonamino, 2-6C alkanoylamino, cyano, nitro, thiol, alkylthio, alkylsulfonyl, alkylaminosulfonyl, 2-6C alkanoyl, aminocarbonyl, N-alkylaminocarbonyl or N,N-amino-carbonyl);

R6,R7 = H or 1-6C alkyl.

Provided that:

(1) when G1 is bicyclic, tricyclic or tetracyclic group, then each ring structure is joined to the next ring structure by a direct bond, -O-, 1-6C alkyl, 1-6C haloalkyl, 1-6C heteroalkyl, 2-6C alkenyl, 2-6C alkynyl, sulfone, CO, NR7CO, CONR7, NR7, S or C(OH) and each ring is fused to the next ring structure; and

(2) when -NR5Y- represents an azacyclic ring and L represents a direct bond, the group G1 is spiro fused to the azacyclic ring.

INDEPENDENT CLAIMS are also included for:

(1) the preparation of (I); and

(2) a pharmaceutical composition comprising compounds of (I) and an adjuvant, diluent or carrier.

ACTIVITY - Antiasthmatic; Respiratory-Gen.; Antiarthritic; Antirheumatic; Osteopathic; Antiarteriosclerotic; Vasotropic; Cytostatic; Antiinflammatory; Cardiant; Hepatotropic; Nephrotropic; Viricide; Gynecological; CNS-Gen.; Neuroprotective; Nootropic; hemostatic.

MECHANISM OF ACTION - Metalloproteinase (MMP) inhibitor (e.g. (MMP12) and (MMP9)); TACE and aggrecanase inhibitors. Isolated enzyme assay was carried out as follows: Recombinant human MMP12 catalytic domain was purified. The purified enzyme was used to monitor inhibitor of activity of MMP12 (50 ng/ml) which was incubated for 60 minutes at room temperature with the synthetic substrate Mac-Pro-Cha-Gly-Nva-His-Ala-Dpa-NH2) in Tris-HCl (RTM; assay buffer) (0.1 M), pH 7.3 containing NaCl (0.1 M), CaCl2 (20 mM), ZnCl (0.020 mM) and Brij 35 (RTM detergent) (0.05 w/v %) in the presence of 5-((4-((2-(trifluoromethyl)pyrimidin-5-yl)ethynyl)-3,6-dihydropyridin-1(2H)-yl)sulfonyl)methyl)-2,4-dihydro-3H-1,2,4-triazol-3-one (test compound). The IC50 value of the test compound was found to be 2.4 nM.

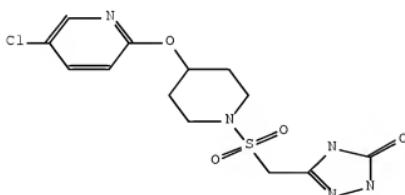
USE - In therapy; in the manufacture of a medicament for the treatment of obstructive airways disease e.g. asthma or chronic obstructive pulmonary disease; for the treatment of disease or condition mediated by metalloproteinases e.g. metalloelastase (MMP12) and gelatinase (MMP9) (all claimed); as metalloproteinase (MMP) inhibitors; as inhibitors of TACE and aggrecanase; as pharmaceuticals. The disease or conditions includes rhinitis, arthritis (such as rheumatoid arthritis and osteoarthritis), atherosclerosis, and restenosis, cancer, invasion and metastasis, diseases involving tissue destruction, loosening of hip joint replacements, periodontal disease, fibrotic disease, infarction and heart disease, liver and renal fibrosis, endometriosis, diseases related to the weakening of the extracellular matrix, heart failure, aortic aneurysms, CNS related diseases such as Alzheimer's disease and multiple sclerosis and hematological disorders, inflammatory diseases.

ADVANTAGE - (I) is inhibitors of metalloproteinases and inhibits MMPs such as MMP12 and MMP9, has beneficial potency, selectivity and pharmacokinetic properties; possess pharmacological activity in animals and thus potentially useful as pharmaceuticals.

AN.S DCR-1175755

CN.S 5-[4-(5-Chloro-pyridin-2-yloxy)-piperidine-1-sulfonylmethyl]-2,4-dihydro-1,2,4-triazol-3-one5-[4-(5-Chloro-pyridin-2-yloxy)-piperidine-1-sulfonylmethyl]-2,4-dihydro-[1,2,4]triazol-3-one

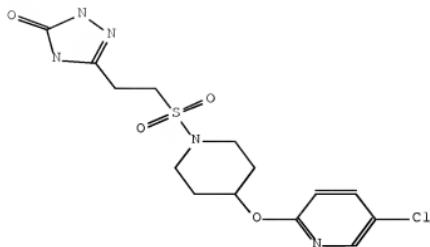
SDCN RAJXZN



AN.S DCR-1175756

CN.S 5-(2-[4-(5-Chloro-pyridin-2-yloxy)-piperidine-1-sulfonyl]-ethyl)-2,4-dihydro-1,2,4-triazol-3-one5-(2-[4-(5-Chloro-pyridin-2-yloxy)-piperidine-1-sulfonyl]-ethyl)-2,4-dihydro-[1,2,4]triazol-3-one

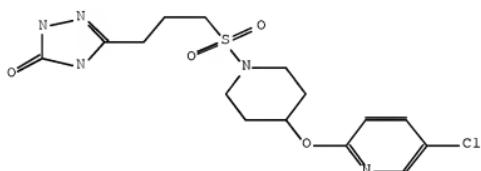
SDCN RAJXZO



AN.S DCR-1175757

CN.S 5-[3-[4-(5-Chloro-pyridin-2-yloxy)-piperidine-1-sulfonyl]-propyl]-2,4-dihydro-1,2,4-triazol-3-one5-[3-[4-(5-Chloro-pyridin-2-yloxy)-piperidine-1-sulfonyl]-propyl]-2,4-dihydro-[1,2,4]triazol-3-one

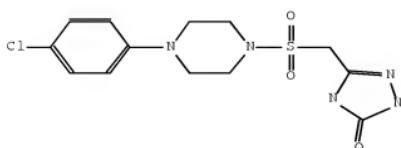
SDCN RAJXZP



AN.S DCR-1175758

CN.S 5-[4-(4-Chloro-phenyl)-piperazine-1-sulfonylmethyl]-2,4-dihydro-1,2,4-triazol-3-one5-[4-(4-Chloro-phenyl)-piperazine-1-sulfonylmethyl]-2,4-dihydro-[1,2,4]triazol-3-one

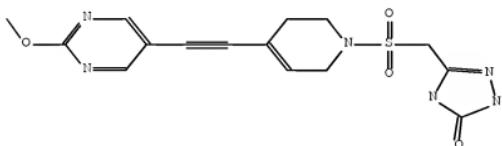
SDCN RAJXZQ



AN.S DCR-1175759

CN.S 5-[4-(2-Methoxy-pyrimidin-5-ylethynyl)-3,6-dihydro-2H-pyridine-1-sulfonylmethyl]-2,4-dihydro-1,2,4-triazol-3-one5-[4-(2-Methoxy-pyrimidin-5-ylethynyl)-3,6-dihydro-2H-pyridine-1-sulfonylmethyl]-2,4-dihydro-[1,2,4]triazol-3-one

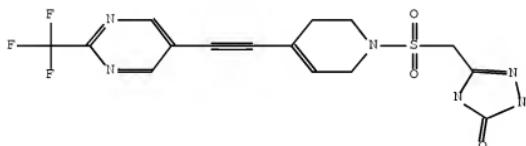
SDCN RAJXZR



AN.S DCR-1175760

CN.S 5-[4-(2-Trifluoromethyl-pyrimidin-5-ylethynyl)-3,6-dihydro-2H-pyridine-1-sulfonylmethyl]-2,4-dihydro-1,2,4-triazol-3-one5-[4-(2-Trifluoromethyl-pyrimidin-5-ylethynyl)-3,6-dihydro-2H-pyridine-1-sulfonylmethyl]-2,4-dihydro-[1,2,4]triazol-3-one

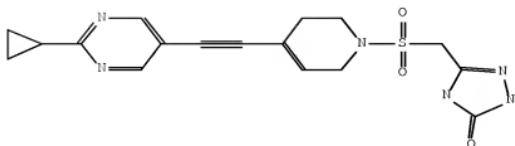
SDCN RAJXZS



AN.S DCR-1175761

CN.S 5-[4-(2-Cyclopropyl-pyrimidin-5-ylethynyl)-3,6-dihydro-2H-pyridine-1-sulfonylmethyl]-2,4-dihydro-1,2,4-triazol-3-one5-[4-(2-Cyclopropyl-pyrimidin-5-ylethynyl)-3,6-dihydro-2H-pyridine-1-sulfonylmethyl]-2,4-dihydro-[1,2,4]triazol-3-one

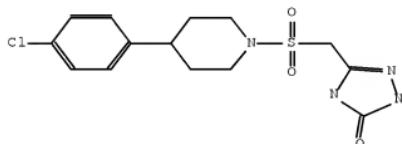
SDCN RAJXZT



AN.S DCR-1175762

CN.S 5-[4-(4-Chloro-phenyl)-piperidine-1-sulfonylmethyl]-2,4-dihydro-1,2,4-triazol-3-one5-[4-(4-Chloro-phenyl)-piperidine-1-sulfonylmethyl]-2,4-dihydro-[1,2,4]triazol-3-one

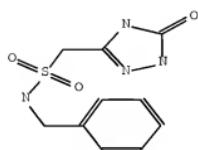
SDCN RAJXZU



AN.S DCR-1175763

CN.S N-Benzyl-C-(5-oxo-4,5-dihydro-1H-1,2,4-triazol-3-yl)-methanesulfonamideN-Benzyl-C-(5-oxo-4,5-dihydro-1H-[1,2,4]triazol-3-yl)-methanesulfonamide

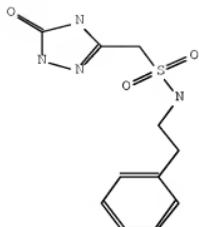
SDCN RAJXZV



AN.S DCR-1175764

CN.S C-(5-Oxo-4,5-dihydro-1H-1,2,4-triazol-3-yl)-N-phenethyl-methanesulfonamideC-(5-Oxo-4,5-dihydro-1H-[1,2,4]triazol-3-yl)-N-phenethyl-methanesulfonamide

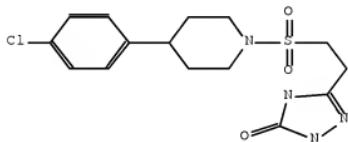
SDCN RAJXZW



AN.S DCR-1175765

CN.S 5-{2-[4-(4-Chloro-phenyl)-piperidine-1-sulfonyl]-ethyl}-2,4-dihydro-1,2,4-triazol-3-one5-{2-[4-(4-Chloro-phenyl)-piperidine-1-sulfonyl]-ethyl}-2,4-dihydro-[1,2,4]triazol-3-one

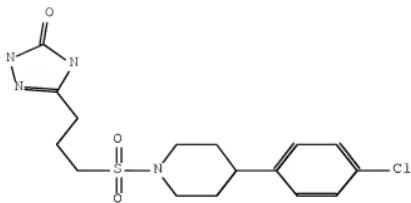
SDCN RAJXZX



AN.S DCR-1175766

CN.S 5-{3-[4-(4-Chloro-phenyl)-piperidine-1-sulfonyl]-propyl}-2,4-dihydro-1,2,4-triazol-3-one5-{3-[4-(4-Chloro-phenyl)-piperidine-1-sulfonyl]-propyl}-2,4-dihydro-[1,2,4]triazol-3-one

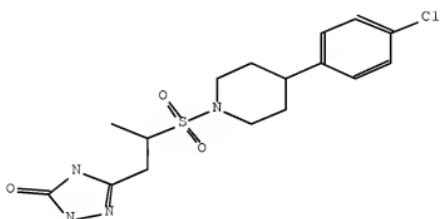
SDCN RAJXZY



AN.S DCR-1175767

CN.S 5-[2-[4-(4-Chloro-phenyl)-piperidine-1-sulfonyl]-propyl]-2,4-dihydro-1,2,4-triazol-3-oneb-[2-[4-(4-Chloro-phenyl)-piperidine-1-sulfonyl]-propyl]-2,4-dihydro-[1,2,4]triazol-3-one

SDCN RAJXZZ



L36 ANSWER 16 OF 17 BABS COPYRIGHT 2009 Elsevier Inf. Sys. on STN
 AN 5704055 BABS [Full-text](#)

TI Synthesis of Azoles and Fused Azoles from α -Arylhydrazoneonitriles

AU Ibrahim, Mohamed Kamal Ahmed; El-Moghayar, Mohamed Riffat Hamza

SO Indian J.Chem.Sect.B (1987), 26(1-12), 832-835

CODEN: IJSBDB

DT Journal

LA English

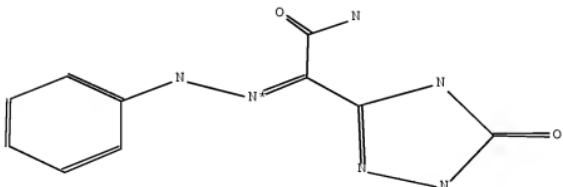
SL English

AB The reaction of α -arylhydrazoneonitriles (I) with phenylhydrazine, hydrazine hydrate, *p*-phenylenediamine, mercaptoacetic acid, and phenyl isothiocyanate gives amidrazone (IIa-c, III), pyrazolone (IVa-c), benzimidazole (X), 4-thiazolone (XIe,d) and 1,2,4-triazinethione (XII) derivatives respectively. Compound IIa reacts with ethyl chloroformate to give amidrazone V which on base-catalyzed cyclization affords the triazolone VI. However, it

reacts with nitrous acid to give the tetrazole VII. The condensation of IVa with ethyl 5a-chloroacetoacetate and IVe with 5a-cyanocinnamonitrile furnishes the pyrazolo<1,5-a>imidazole (VIII) and pyrano<3,2-b>pyrazole (IX) derivatives respectively.

L36 ANSWER 17 OF 17 BEILSTEIN COPYRIGHT 2009 Elsevier Inf. Sys. on STN

Beilstein Records (BRN): 5574184
 Beilstein Pref. RN (BPR): 115998-45-3
 CAS Reg. No. (RN): 115998-45-3
 Chemical Name (CN): 5-<phenylhydrazone-(carbamoyl)-methyl>-1,2,
 ,4-triazol-3-one
 Autonom Name (AUN): 2-(5-oxo-4,5-dihydro-1H-<1,2,4>triazol-3-y
 l)-2-(phenyl-hydrazone)-acetamide
 Molec. Formula (MF): C10 H10 N6 O2
 Molecular Weight (MW): 246.23
 Lawson Number (LN): 30257, 16435
 Compound Type (CTYPE): heterocyclic
 Constitution ID (CONSID): 4886893
 Beilstein Citation (BSO): 6-26
 Entry Date (DED): 1993/02/12
 Update Date (DUPD): 1994/02/18



Field Availability:

| Code | Name | Occurrence |
|-------|------------------------|------------|
| ===== | | |
| BRN | Beilstein Records | 1 |
| BPR | Beilstein Preferred RN | 1 |
| RN | CAS Registry Number | 1 |
| CN | Chemical Name | 1 |

| | | |
|--------|----------------------------|---|
| AUN | Autonomname | 1 |
| MF | Molecular Formula | 1 |
| FW | Formular Weight | 1 |
| LN | Lawson Number | 2 |
| CTYPE | Compound Type | 1 |
| CONSID | Constitution ID | 1 |
| BSO | Beilstein Citation | 1 |
| DED | Entry Date | 1 |
| DUPD | Update Date | 1 |
| IR | Infrared Spectrum | 1 |
| MP | Melting Point | 1 |
| NMR | Nuclear Magnetic Resonance | 1 |

This substance also occurs in Reaction Documents:

| Code | Name | Occurrence |
|-------|-------------------------------|------------|
| RX | Reaction Documents | 1 |
| RXPRO | Substance is Reaction Product | 1 |

All References:

ALLREF

1. Ibrahim, Mohamed Kamal Ahmed; El-Moghayar, Mohamed Riffat Hamza, Indian J.Chem.Sect.B, CODEN: IJSBDB, 26(1-12), <1987>, 832-835; BABS-5704055

Search History

L23 STRUCTURE UPLOADED
L24 0 SEA SSS SAM L23
L25 0 SEA SSS FUL L23

FILE 'MARPAT' ENTERED AT 14:19:06 ON 02 OCT 2009
L26 STRUCTURE UPLOADED
L27 6 SEA SSS SAM L26
L28 93 SEA SSS FUL L26
L29 STRUCTURE UPLOADED
L30 6 SEA SUB=L28 SSS SAM L29
L31 87 SEA SUB=L28 SSS FUL L29

FILE 'MARPAT' ENTERED AT 14:52:40 ON 02 OCT 2009
L32 STRUCTURE UPLOADED

FILE 'HCAPLUS, WPIX' ENTERED AT 15:05:48 ON 02 OCT 2009
L33 1 DUP REM L11 L15 (1 DUPLICATE REMOVED)

FILE 'HCAPLUS' ENTERED AT 15:06:05 ON 02 OCT 2009
L34 14 SEA SPE=ON ABB=ON PLU=ON L8 NOT L11

FILE 'WPIX' ENTERED AT 15:06:20 ON 02 OCT 2009
L35 0 SEA SPE=ON ABB=ON PLU=ON L14 NOT L15

FILE 'HCAPLUS, WPIX, BABS, BEILSTEIN' ENTERED AT 15:07:19 ON 02 OCT 2009
L36 17 DUP REM L34 L14 L19 L17 (0 DUPLICATES REMOVED)